This listing of claims as required by 37 C.F.R. 1.121 will replace all prior versions and listing of claims in the application:

This listing of claims will replace all prior versions and listings of claims in the application:

Listing of Claims:

1. (Original) A method of inhibiting HDAC-4 activity in a cell, comprising contacting the cell with an antisense oligonucleotide complementary to a region of RNA that encodes a portion of HDAC-4, whereby HDAC-4 activity is inhibited.

2. (Original) The method according to claim 1, wherein the cell is contacted with an HDAC-4 antisense oligonucleotide that is a chimeric oligonucleotide.

3. (Original) The method according to claim 1, wherein the cell is contacted with an HDAC-4 antisense oligonucleotide that is a hybrid oligonucleotide.

4. (Original) The method according to claim 1, wherein the antisense oligonucleotide has a nucleotide sequence of from about 13 to about 35 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:4.

5. (Currently Amended) The method according to claim 1, wherein the antisense oligonucleotide has a nucleotide sequence of from about 15 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:4~~.~~

~~The method according to claim 1, wherein the antisense oligonucleotide has a nucleotide sequence of from about 20 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:4.~~

6. (Original) The method according to claim 1, wherein the cell is contacted with an HDAC-4 antisense oligonucleotide that is SEQ ID NO:11.

7. (Currently Amended) The ~~composition~~ method according to claim 2, wherein the antisense oligonucleotide has a nucleotide sequence of from about 20 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:4.

8. (Currently Amended) The ~~composition~~ method according to claim 2, wherein the antisense oligonucleotide is SEQ ID NO:11.

9. (Currently Amended) The ~~composition~~ method according to claim 2, wherein the antisense oligonucleotide has one or more phosphorothioate internucleoside linkages.

10. (Currently Amended) The ~~composition~~ method according to claim 9, wherein the antisense oligonucleotide further comprises a length of 20-26 nucleotides.

11. (Currently Amended) The ~~composition~~ method according to claim 10, wherein the oligonucleotide is modified such that the terminal four nucleotides at the 5' end of the oligonucleotide and the terminal four nucleotides at the 3' end of the oligonucleotide each have 2' -O- methyl groups attached to their sugar residues.

12. (Currently Amended) ~~The composition according to claim 1, wherein the agent is a small molecule inhibitor of HDAC-4.~~A method of

inhibiting HDAC-4 activity in a cell, comprising contacting the cell with a small molecule inhibitor of HDAC-4.

13. (Currently Amended) The ~~composition~~ method according to claim 12, wherein the structure of the small molecule inhibitor is selected from the group consisting of:

(a) Cy-CH(OMe)-Y^1-C(O)-NH-Z (1)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; Y^1 is a C_4 - C_6 alkylene, wherein said alkylene may be optionally substituted and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR^1, R^1 being alkyl, acyl or hydrogen; S; S(O); or $S(O)_2$; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl and -O-M, M being H or a pharmaceutically acceptable cation, wherein the anilinyl or pyridyl or thiadiazolyl may be optionally substituted;

(b) Cy-Y^2-C(O)-NH-Z (2)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; Y^2 is C_5 - C_7 alkylene, wherein said alkylene may be optionally substituted and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR^1, R^1 being alkyl, acyl or hydrogen; S; S(O); or $S(O)_2$; and Z is anilinyl or pyridyl, or thiadiazolyl, any of which may be optionally substituted;

(c) Cy-B-Y^3-C(O)-NH-Z (3)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; B is selected from the group consisting of

-CH(OMe), ketone and methylene; Y^3 is a C_4 - C_6 alkylene, wherein said alkylene may be optionally substituted and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR^1, R^1 being alkyl, acyl or hydrogen; S; S(O); or $S(O)_2$; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl and -O-M, M being H or a pharmaceutically acceptable cation, wherein the anilinyl or pyridyl or thiadiazolyl may be optionally substituted;

(d) $Cy–L^1–Ar–Y^1–C(O)–NH-Z$ (4)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; L^1 is $-(CH_2)_m-W-$, where m is 0, 1, 2, 3, or 4, and W is selected from the group consisting of –C(O)NH–, –$S(O)_2NH$–, -NHC(O)-, $-NHS(O)_2-$, and -NH-C(O)-NH-; Ar is arylene, wherein said arylene optionally may be additionally substituted and optionally may be fused to an aryl or heteroaryl ring, or to a saturated or partially unsaturated cycloalkyl or heterocyclic ring, any of which may be optionally substituted; Y^1 is a chemical bond or a straight- or branched-chain saturated alkylene, wherein said alkylene may be optionally substituted; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl, and -O-M, M being H or a pharmaceutically acceptable cation; provided that when L^1 is –C(O)NH–, Y^1 is $–(CH_2)_n–$, n being 1, 2, or 3, and Z is -O-M, then Cy is not aminophenyl, dimethylaminophenyl, or hydroxyphenyl; and further provided that when L^1 is -C(O)NH- and Z is pyridyl, then Cy is not substituted indolinyl;

(e) $Cy–L^2–Ar–Y^2–C(O)NH-Z$ (5)

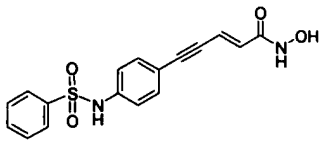
wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted, provided that Cy is not a

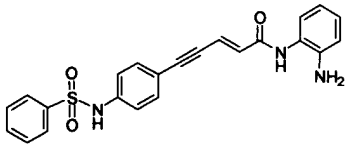
(spirocycloalkyl)heterocyclyl; L^2 is C_1-C_6 saturated alkylene or C_2-C_6 alkenylene, wherein the alkylene or alkenylene optionally may be substituted, provided that L^2 is not –C(O)–, and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR', R' being alkyl, acyl, or hydrogen; S; S(O); or $S(O)_2$; Ar is arylene, wherein said arylene optionally may be additionally substituted and optionally may be fused to an aryl or heteroaryl ring, or to a saturated or partially unsaturated cycloalkyl or heterocyclic ring, any of which may be optionally substituted; and Y^2 is a chemical bond or a straight- or branched-chain saturated alkylene, which may be optionally substituted, provided that the alkylene is not substituted with a substituent of the formula –C(O)R wherein R comprises an α-amino acyl moiety; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl, and -O-M, M being H or a pharmaceutically acceptable cation; provided that when the carbon atom to which Cy is attached is oxo substituted, then Cy and Z are not both pyridyl;

(f) $Cy–L^3–Ar–Y^3–C(O)NH-Z$ (6)

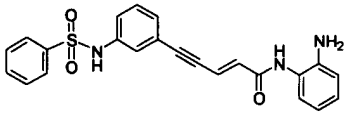
wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted, provided that Cy is not a (spirocycloalkyl)heterocyclyl; L^3 is selected from the group consisting of (a) $-(CH_2)_m-W-$, where m is 0, 1, 2, 3, or 4, and W is selected from the group consisting of –C(O)NH–, $–S(O)_2NH–$, -NHC(O)-, $-NHS(O)_2-$, and -NH-C(O)-NH-; and (b) C_1-C_6 alkylene or C_2-C_6 alkenylene, wherein the alkylene or alkenylene optionally may be substituted, provided that L^3 is not –C(O)–, and wherein one of the carbon atoms of the alkylene optionally may be replaced by O; NR', R' being alkyl, acyl, or hydrogen; S; S(O); or $S(O)_2$; Ar is arylene, wherein said arylene optionally may be additionally substituted and optionally may be fused to an aryl or

heteroaryl ring, or to a saturated or partially unsaturated cycloalkyl or heterocyclic ring, any of which may be optionally substituted; and Y^3 is C_2 alkenylene or C_2 alkynylene, wherein one or both carbon atoms of the alkenylene optionally may be substituted with alkyl, aryl, alkaryl, or aralkyl; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl, and -O-M, M being H or a pharmaceutically acceptable cation; provided that when Cy is unsubstituted phenyl, Ar is not phenyl wherein L^3 and Y^3 are oriented *ortho* or *meta* to each other;

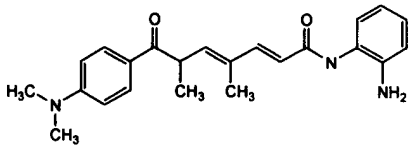
~~(e)~~(g)  (7)

~~(f)~~(h)  (8)

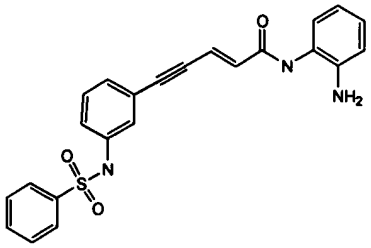
and

~~(g)~~(i)  (9).

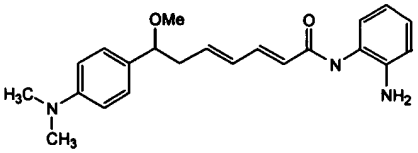
14. (Currently Amended) The ~~composition~~ method according to claim 13, wherein the small molecule inhibitor is selected from the group consisting of:

(a) 

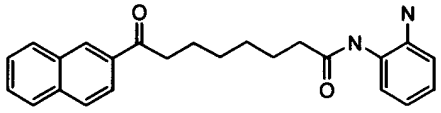
(b)



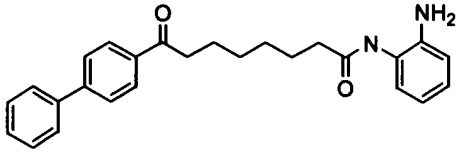
(c)



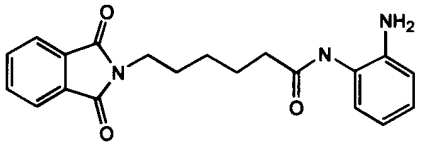
(d)



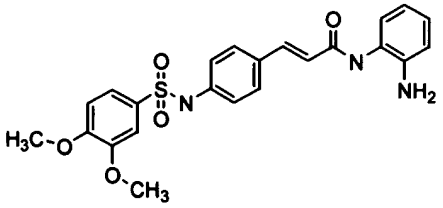
(e)



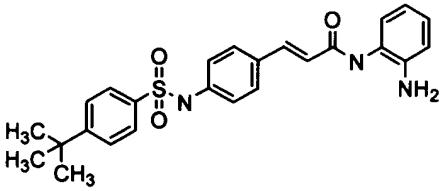
(f)



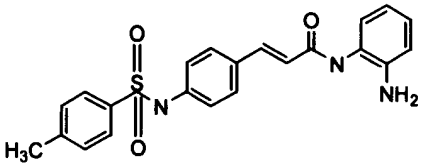
(g)



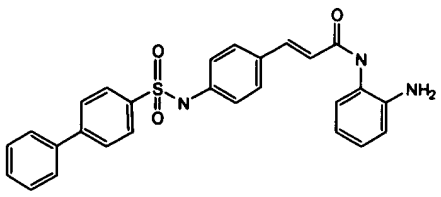
(h)



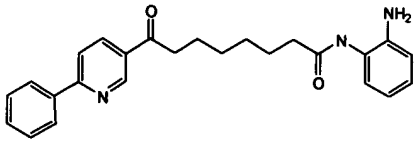
(i)



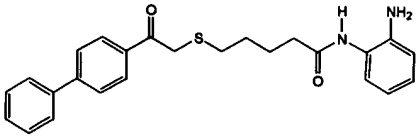
(j)



(k)



(l)



15. (Original) A method for inhibiting HDAC-4 activity in a cell, comprising contacting the cell with a specific inhibitor of HDAC-4, whereby HDAC-4 activity is inhibited.

16. (Original) The method according to claim 15, wherein the cell is contacted with a specific inhibitor of HDAC-4 activity selected from the group consisting of:

(a) an antisense oligonucleotide complementary to a region of RNA that encodes a portion of HDAC-4, and

(b) a small molecule inhibitor of HDAC-4.

17. (Original) The method according to claim 16, wherein the specific inhibitor is an antisense oligonucleotide complementary to a region of RNA that encodes a portion of HDAC-4.

18. (Original) The method according to claim 17, wherein the cell is contacted with an HDAC-4 antisense oligonucleotide that is a chimeric oligonucleotide.

19. (Original) The method according to claim 17, wherein the cell is contacted with an HDAC-4 antisense oligonucleotide that is a hybrid oligonucleotide.

20. (Original) The method according to claim 17, wherein the cell is contacted with an HDAC-4 antisense oligonucleotide that has a nucleotide sequence length of from about 13 to about 35 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:4.

21. (Original) The method according to claim 17, wherein the cell is contacted with an HDAC-4 antisense oligonucleotide that has a nucleotide sequence length of from about 15 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:4.

22. (Original) The method according to claim 17, wherein the cell is contacted with an HDAC-4 antisense oligonucleotide that has a nucleotide sequence length of from about 20 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:4.

23. (Original) The method according to claim 17, wherein the cell is contacted with an DHAC-4 antisense oligonucleotide that is SED ID NO:11.

24. (Original) The method according to claim 16 wherein the small molecule inhibitor of HDAC-4 has a structure selected form the group consisting of:

(a) Cy-CH(OMe)-Y^1-C(O)-NH-Z (1)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; Y^1 is a C_4 - C_6 alkylene, wherein said alkylene may be optionally substituted and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom

moiety selected from the group consisting of O; NR^1, R^1 being alkyl, acyl or hydrogen; S; S(O); or $S(O)_2$; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl and -O-M, M being H or a pharmaceutically acceptable cation, wherein the anilinyl or pyridyl or thiadiazolyl may be optionally substituted;

(b) $Cy\text{-}Y^2\text{-}C(O)\text{-}NH\text{-}Z$ (2)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; Y^2 is C_5 - C_7 alkylene, wherein said alkylene may be optionally substituted and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR^1, R^1 being alkyl, acyl or hydrogen; S; S(O); or $S(O)_2$; and Z is anilinyl or pyridyl, or thiadiazolyl, any of which may be optionally substituted;

(c) $Cy\text{-}B\text{-}Y^3\text{-}C(O)\text{-}NH\text{-}Z$ (3)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; B is selected from the group consisting of -CH(OMe), ketone and methylene; Y^3 is a C_4 - C_6 alkylene, wherein said alkylene may be optionally substituted and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR^1, R^1 being alkyl, acyl or hydrogen; S; S(O); or $S(O)_2$; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl and -O-M, M being H or a pharmaceutically acceptable cation, wherein the anilinyl or pyridyl or thiadiazolyl may be optionally substituted;

(d) $Cy\text{–}L^1\text{–}Ar\text{–}Y^1\text{–}C(O)\text{–}NH\text{-}Z$ (4)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; L^1 is $-(CH_2)_m\text{-}W\text{-}$, where m is 0, 1, 2, 3, or

4, and W is selected from the group consisting of –C(O)NH–, –$S(O)_2NH$–, -NHC(O)-, -$NHS(O)_2$-, and -NH-C(O)-NH-; Ar is arylene, wherein said arylene optionally may be additionally substituted and optionally may be fused to an aryl or heteroaryl ring, or to a saturated or partially unsaturated cycloalkyl or heterocyclic ring, any of which may be optionally substituted; Y^1 is a chemical bond or a straight- or branched-chain saturated alkylene, wherein said alkylene may be optionally substituted; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl, and -O-M, M being H or a pharmaceutically acceptable cation; provided that when L^1 is –C(O)NH, Y^1 is –$(CH_2)_n$–, n being 1, 2, or 3, and Z is -O-M, then Cy is not aminophenyl, dimethylaminophenyl, or hydroxyphenyl; and further provided that when L^1 is -C(O)NH- and Z is pyridyl, then Cy is not substituted indolinyl;

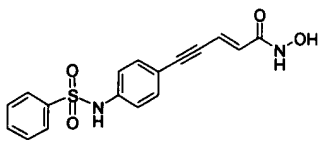
(e) $Cy–L^2–Ar–Y^2–C(O)NH-Z$ (5)

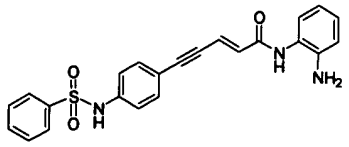
wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted, provided that Cy is not a (spirocycloalkyl)heterocyclyl; L^2 is C_1-C_6 saturated alkylene or C_2-C_6 alkenylene, wherein the alkylene or alkenylene optionally may be substituted, provided that L^2 is not –C(O)–, and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR', R' being alkyl, acyl, or hydrogen; S; S(O); or $S(O)_2$; Ar is arylene, wherein said arylene optionally may be additionally substituted and optionally may be fused to an aryl or heteroaryl ring, or to a saturated or partially unsaturated cycloalkyl or heterocyclic ring, any of which may be optionally substituted; and Y^2 is a chemical bond or a straight- or branched-chain saturated alkylene, which may be optionally substituted, provided that the alkylene is not substituted with a substituent of the

formula –C(O)R wherein R comprises an α-amino acyl moiety; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl, and -O-M, M being H or a pharmaceutically acceptable cation; provided that when the carbon atom to which Cy is attached is oxo substituted, then Cy and Z are not both pyridyl;

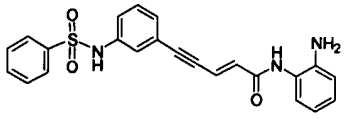
(f) $Cy–L^3–Ar–Y^3–C(O)NH-Z$ (6)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted, provided that Cy is not a (spirocycloalkyl)heterocyclyl; L^3 is selected from the group consisting of (a) $-(CH_2)_m-W-$, where m is 0, 1, 2, 3, or 4, and W is selected from the group consisting of –C(O)NH–, $–S(O)_2NH–$, -NHC(O)-, $-NHS(O)_2-$, and -NH-C(O)-NH-; and (b) C_1-C_6 alkylene or C_2-C_6 alkenylene, wherein the alkylene or alkenylene optionally may be substituted, provided that L^3 is not –C(O)–, and wherein one of the carbon atoms of the alkylene optionally may be replaced by O; NR', R' being alkyl, acyl, or hydrogen; S; S(O); or $S(O)_2$; Ar is arylene, wherein said arylene optionally may be additionally substituted and optionally may be fused to an aryl or heteroaryl ring, or to a saturated or partially unsaturated cycloalkyl or heterocyclic ring, any of which may be optionally substituted; and Y^3 is C_2 alkenylene or C_2 alkynylene, wherein one or both carbon atoms of the alkenylene optionally may be substituted with alkyl, aryl, alkaryl, or aralkyl; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl, and -O-M, M being H or a pharmaceutically acceptable cation; provided that when Cy is unsubstituted phenyl, Ar is not phenyl wherein L^3 and Y^3 are oriented *ortho* or *meta* to each other;

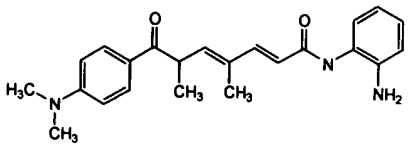
~~(e)~~(g)  (7)

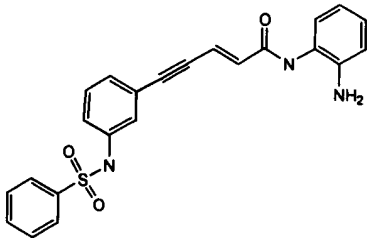
~~(f)~~(h)  (8)

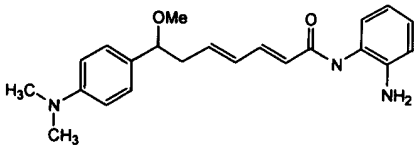
and

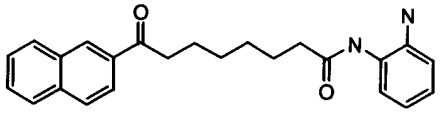
~~(g)~~(i)  (9).

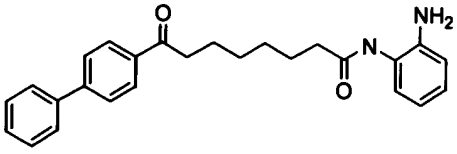
25. (Original) The method according to claim 24, wherein the small molecule inhibitor is selected from the group consisting of:

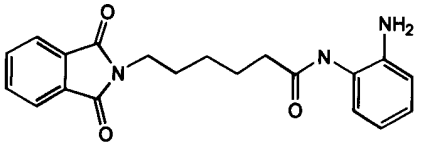
(a) 

(b) 

(c) 

(d) 

(e) 

(f) 

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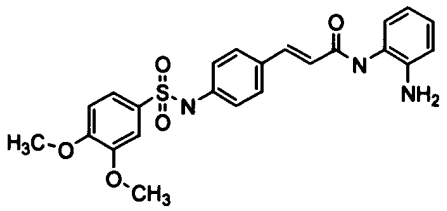
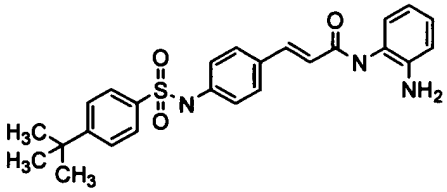


(h)



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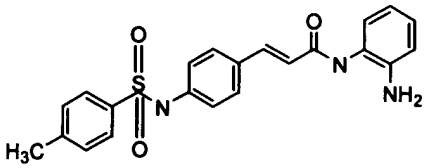
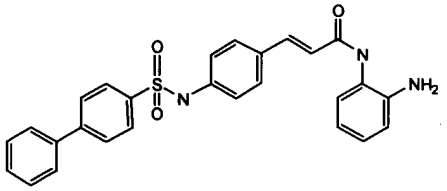
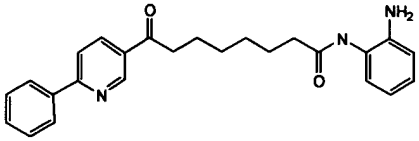


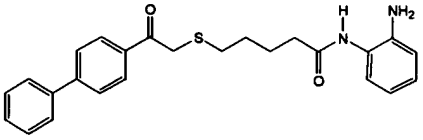
(j)



(k)



(l)



26. (Original) The method according to claim 15, wherein inhibition of HDAC-4 activity in the contacted cell further leads to an inhibition of cell proliferation in the contacted cell.

27. (Original) The method according to claim 15, wherein inhibition of HDAC-4 activity in the contacted cell further leads to growth retardation of the contacted cell.

28. (Original) A method according to claim 15, wherein inhibition of HDAC-4 activity in the contacted cell further leads to growth arrest of the contacted cell.

29. (Original) The method according to claim 15, wherein the inhibition of DHAC-4 activity in the contacted cell further leads to programmed cell death of the contacted cell.

30. (Original) The method according to claim 26, wherein inhibition of HDAC-4 activity in the contacted cell further leads to necrotic cell death of the contacted cell.

31. (Original) A method for inhibiting neoplastic cell proliferation in an animal, comprising administering to an animal having at least one neoplastic cell present in its body a therapeutically effective amount of at least one specific inhibitor of HDAC-4, whereby neoplastic cell proliferation is inhibited in the animal.

32. (Original) The method according to claim 31, wherein the animal is administered a specific inhibitor of HDAC-4 selected from the group consisting of:

(a) an antisense oligonucleotide complementary to a region of RNA that encodes a portion of HDAC-4, and

(b) a small molecule inhibitor.

33. (Original) The method according to claim 32, wherein the animal is administered a therapeutically effective amount of an antisense oligonucleotide complementary to a region of RNA that encodes a portion of HDAC-4, whereby neoplastic cell proliferation is inhibited in the animal.

34. (Original) The method according to claim 33, wherein the animal is administered a chimeric HDAC-4 antisense oligonucleotide.

35. (Original) The method according to claim 33, wherein the animal is administered a hybrid HDAC-4 antisense oligonucleotide.

36. (Original) The method according to claim 33, wherein the animal is administered an HDAC-4 antisense oligonucleotide having a nucleotide sequence of from about 13 to about 35 nucleotides which is selected form the nucleotide sequence of SED IS NO:4.

37. (Original) The method according to claim 32, wherein the animal is administered an HDAC-4 antisense oligonucleotide having a nucleotide sequence of form about 15 to about 26 nucleotides which is selected from the nucleotide sequence of SED IS NO:4.

38. (Original) The method according to claim 32, wherein the cell is contacted with an HDAC-4 antisense oligonucleotide that has a nucleotide sequence length of from about 20 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:4.

39. (Original) The method according to claim 32, wherein the animal is administered an HDAC-4 antisense oligonucleotide that is SEQ ID NO:11.

40. (Currently Amended) The method according to claim 32, wherein a specific inhibitor is a small molecule inhibitor of HDAC-4 having a structure selected from the group consisting of:

(a) $Cy\text{-}CH(OMe)\text{-}Y^1\text{-}C(O)\text{-}NH\text{-}Z$ (1)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; Y^1 is a C_4 - C_6 alkylene, wherein said alkylene may be optionally substituted and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR^1, R^1 being alkyl, acyl or hydrogen; S; S(O); or $S(O)_2$; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl and -O-M, M being H or a pharmaceutically acceptable cation, wherein the anilinyl or pyridyl or thiadiazolyl may be optionally substituted;

(b) $Cy\text{-}Y^2\text{-}C(O)\text{-}NH\text{-}Z$ (2)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; Y^2 is C_5 - C_7 alkylene, wherein said alkylene may be optionally substituted and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR^1, R^1 being alkyl, acyl or hydrogen; S; S(O); or $S(O)_2$; and Z is anilinyl or pyridyl, or thiadiazolyl, any of which may be optionally substituted;

(c) $Cy\text{-}B\text{-}Y^3\text{-}C(O)\text{-}NH\text{-}Z$ (3)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; B is selected from the group consisting of -CH(OMe), ketone and methylene; Y^3 is a C_4 - C_6 alkylene, wherein said alkylene may be optionally substituted and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR^1, R^1 being alkyl, acyl or hydrogen; S; S(O); or $S(O)_2$; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl and -O-M, M being H or a pharmaceutically acceptable cation, wherein the anilinyl or pyridyl or thiadiazolyl may be optionally substituted;

(d) $Cy\text{–}L^1\text{–}Ar\text{–}Y^1\text{–}C(O)\text{–}NH\text{-}Z$ (4)

wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted; L^1 is $-(CH_2)_m\text{-}W\text{-}$, where m is 0, 1, 2, 3, or 4, and W is selected from the group consisting of –C(O)NH–, $–S(O)_2NH–$, -NHC(O)-, $-NHS(O)_2$-, and -NH-C(O)-NH-; Ar is arylene, wherein said arylene optionally may be additionally substituted and optionally may be fused to an aryl or heteroaryl ring, or to a saturated or partially unsaturated cycloalkyl or heterocyclic ring, any of which may be optionally substituted; Y^1 is a chemical bond or a straight- or branched-chain saturated alkylene, wherein said alkylene

may be optionally substituted; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl, and -O-M, M being H or a pharmaceutically acceptable cation; provided that when L^1 is –C(O)NH–, Y^1 is $-(CH_2)_n-$, n being 1, 2, or 3, and Z is -O-M, then Cy is not aminophenyl, dimethylaminophenyl, or hydroxyphenyl; and further provided that when L^1 is -C(O)NH- and Z is pyridyl, then Cy is not substituted indolinyl;

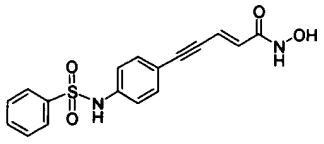
(e) $Cy–L^2–Ar–Y^2–C(O)NH-Z$ (5)

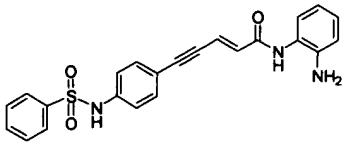
wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted, provided that Cy is not a (spirocycloalkyl)heterocyclyl; L^2 is C_1-C_6 saturated alkylene or C_2-C_6 alkenylene, wherein the alkylene or alkenylene optionally may be substituted, provided that L^2 is not –C(O)–, and wherein one of the carbon atoms of the alkylene optionally may be replaced by a heteroatom moiety selected from the group consisting of O; NR', R' being alkyl, acyl, or hydrogen; S; S(O); or $S(O)_2$; Ar is arylene, wherein said arylene optionally may be additionally substituted and optionally may be fused to an aryl or heteroaryl ring, or to a saturated or partially unsaturated cycloalkyl or heterocyclic ring, any of which may be optionally substituted; and Y^2 is a chemical bond or a straight- or branched-chain saturated alkylene, which may be optionally substituted, provided that the alkylene is not substituted with a substituent of the formula –C(O)R wherein R comprises an α-amino acyl moiety; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl, and -O-M, M being H or a pharmaceutically acceptable cation; provided that when the carbon atom to which Cy is attached is oxo substituted, then Cy and Z are not both pyridyl;

(f) $Cy–L^3–Ar–Y^3–C(O)NH-Z$ (6)

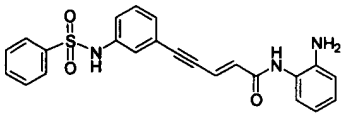
wherein Cy is cycloalkyl, aryl, heteroaryl, or heterocyclyl, any of which may be optionally substituted, provided that Cy is not a (spirocycloalkyl)heterocyclyl; L^3 is selected from the group consisting of (a) $-(CH_2)_m-W-$, where m is 0, 1, 2, 3,

or 4, and W is selected from the group consisting of –C(O)NH–, $-S(O)_2NH-$, -NHC(O)-, $-NHS(O)_2-$, and -NH-C(O)-NH-; and (b) C_1-C_6 alkylene or C_2-C_6 alkenylene, wherein the alkylene or alkenylene optionally may be substituted, provided that L^3 is not –C(O)–, and wherein one of the carbon atoms of the alkylene optionally may be replaced by O; NR', R' being alkyl, acyl, or hydrogen; S; S(O); or $S(O)_2$; Ar is arylene, wherein said arylene optionally may be additionally substituted and optionally may be fused to an aryl or heteroaryl ring, or to a saturated or partially unsaturated cycloalkyl or heterocyclic ring, any of which may be optionally substituted; and Y^3 is C_2 alkenylene or C_2 alkynylene, wherein one or both carbon atoms of the alkenylene optionally may be substituted with alkyl, aryl, alkaryl, or aralkyl; and Z is selected from the group consisting of anilinyl, pyridyl, thiadiazolyl, and -O-M, M being H or a pharmaceutically acceptable cation; provided that when Cy is unsubstituted phenyl, Ar is not phenyl wherein L^3 and Y^3 are oriented *ortho* or *meta* to each other;

~~(e)~~(g)  (7)

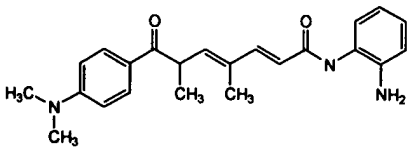
~~(f)~~(h)  (8)

and

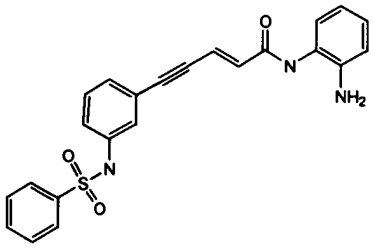
~~(g)~~(i)  (9).

41. (Original) The method according to claim 40, wherein the small molecule inhibitor is selected from the group consisting of:

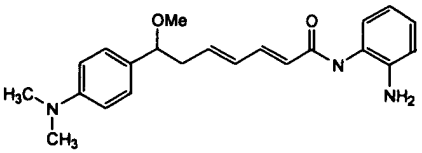
(a)



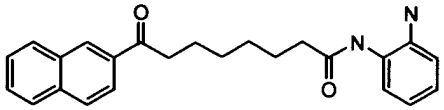
(b)



(c)



(d)



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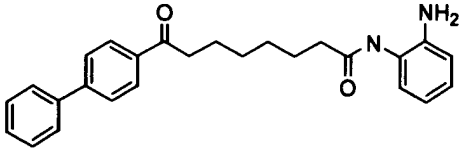

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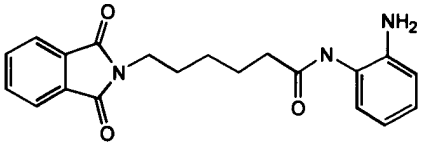

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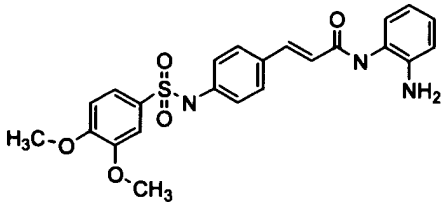

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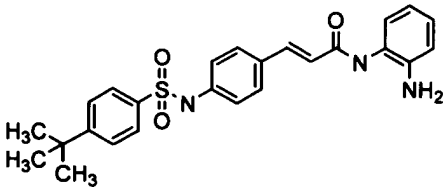

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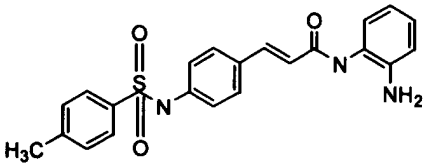

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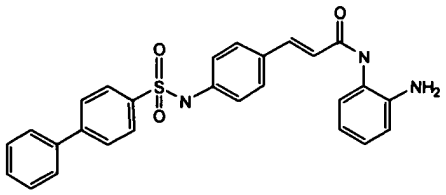
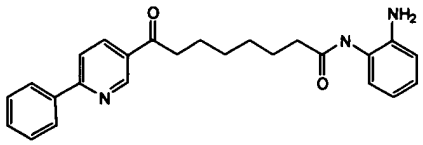


(k)



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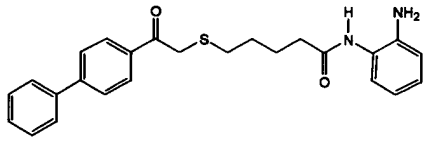

42. (Original) The method according to claim 32, further comprising administering to an animal a therapeutically effective amount of an antisense oligonucleotide complementary to a region of RNA that encodes a portion of HDAC-1.

43. (Original) The method according to claim 42, wherein the animal is administered a chimeric HDAC-1 antisense oligonucleotide.

44. (Original) The method according to claim 42, wherein the animal is administered a hybrid HDAC-1 antisense oligonucleotide.

45. (Original) The method according to claim 42, wherein the animal is administered an HDAC-1 antisense oligonucleotide having a nucleotide sequence from about 13 to about 35 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:2.

46. (Original) The method according to claim 42, wherein the animal is administered an HDAC-1 antisense oligonucleotide having a nucleotide sequence of from about 15 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:2.

47. (Original) The method according to claim 42, wherein the animal is administered an HDAC-1 antisense oligonucleotide having a nucleotide sequence of from about 20 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:2.

48. (Original) The method according to claim 42, wherein the animal is administered an HDAC-1 antisense oligonucleotide that is SEQ ID NO:5

49. (Previously Added) The method according to claim 1, wherein the antisense oligonucleotide has a nucleotide sequence of from about 20 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID No.: 4.

50. (New) The method according to claim 1, wherein the antisense oligonucleotide has a nucleotide sequence of from about 20 to about 26 nucleotides which is selected from the nucleotide sequence of SEQ ID NO:4.

CONCLUSION

If the Examiner is of the opinion that a telephone conference would expedite prosecution of the captioned application, the Examiner is encouraged to contact Applicants' undersigned representative.

Respectfully submitted,

Dated: 9/11/03

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